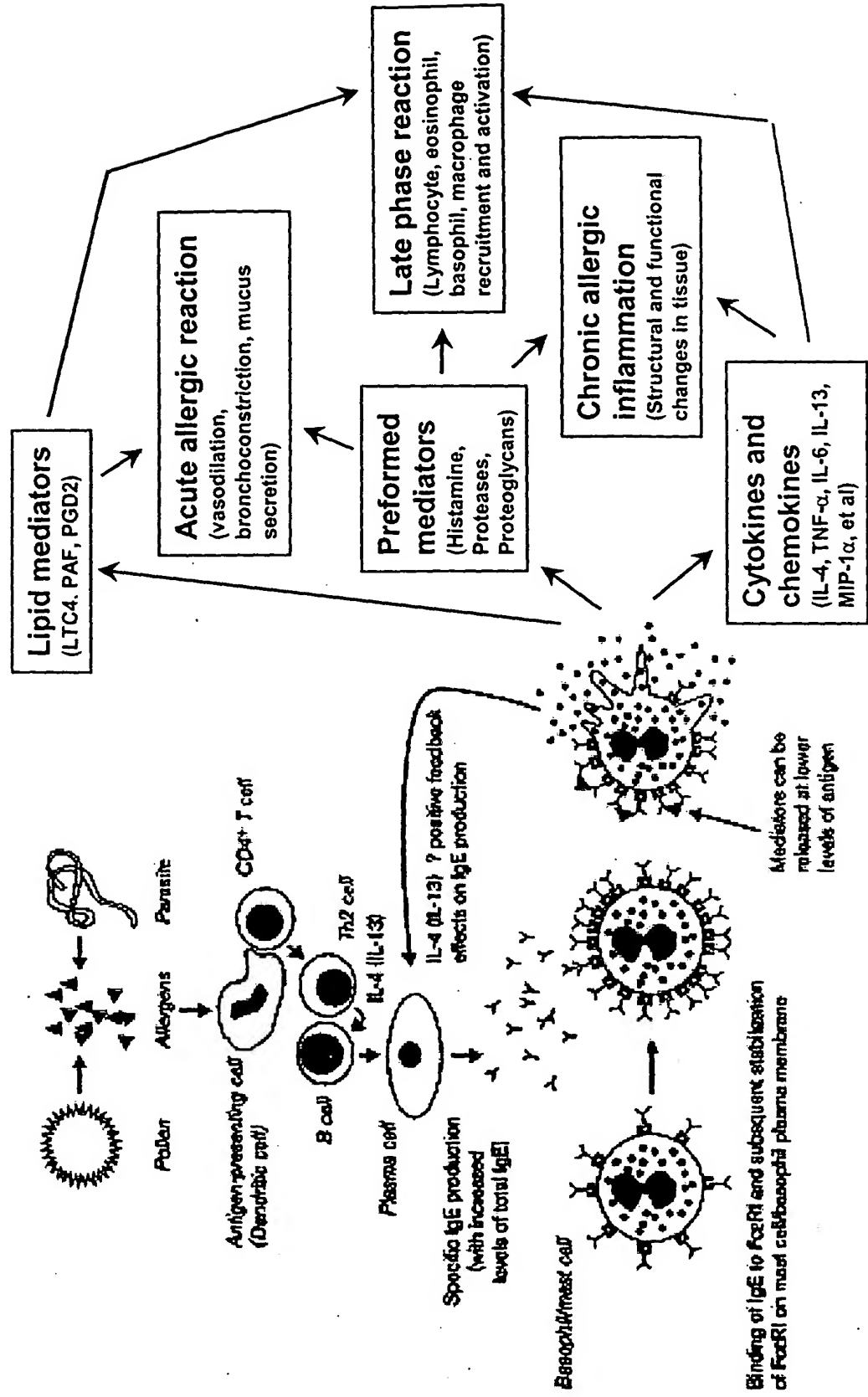


FIG. 1



Mast Cell FceR1 Signaling Pathway

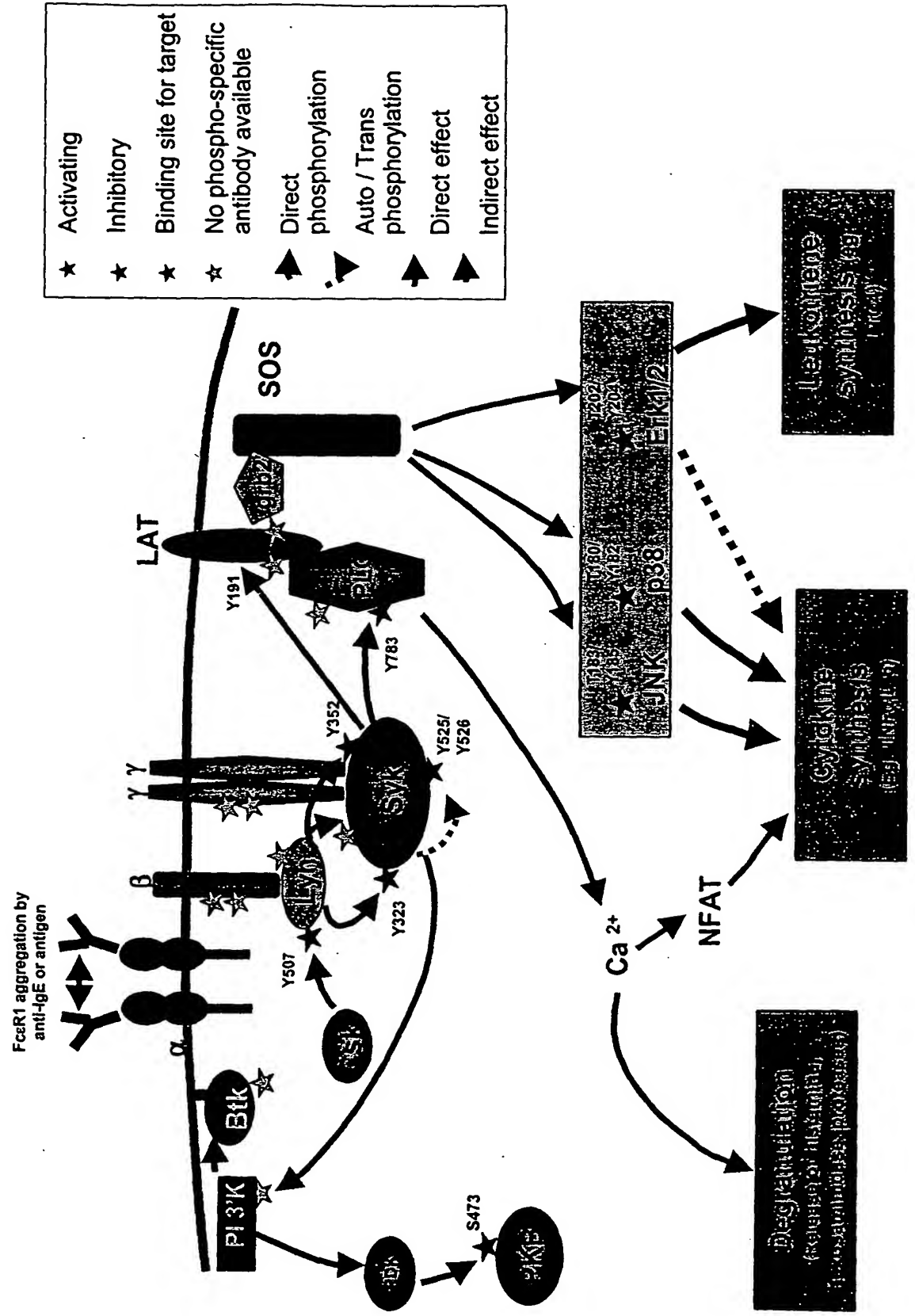


FIG. 3

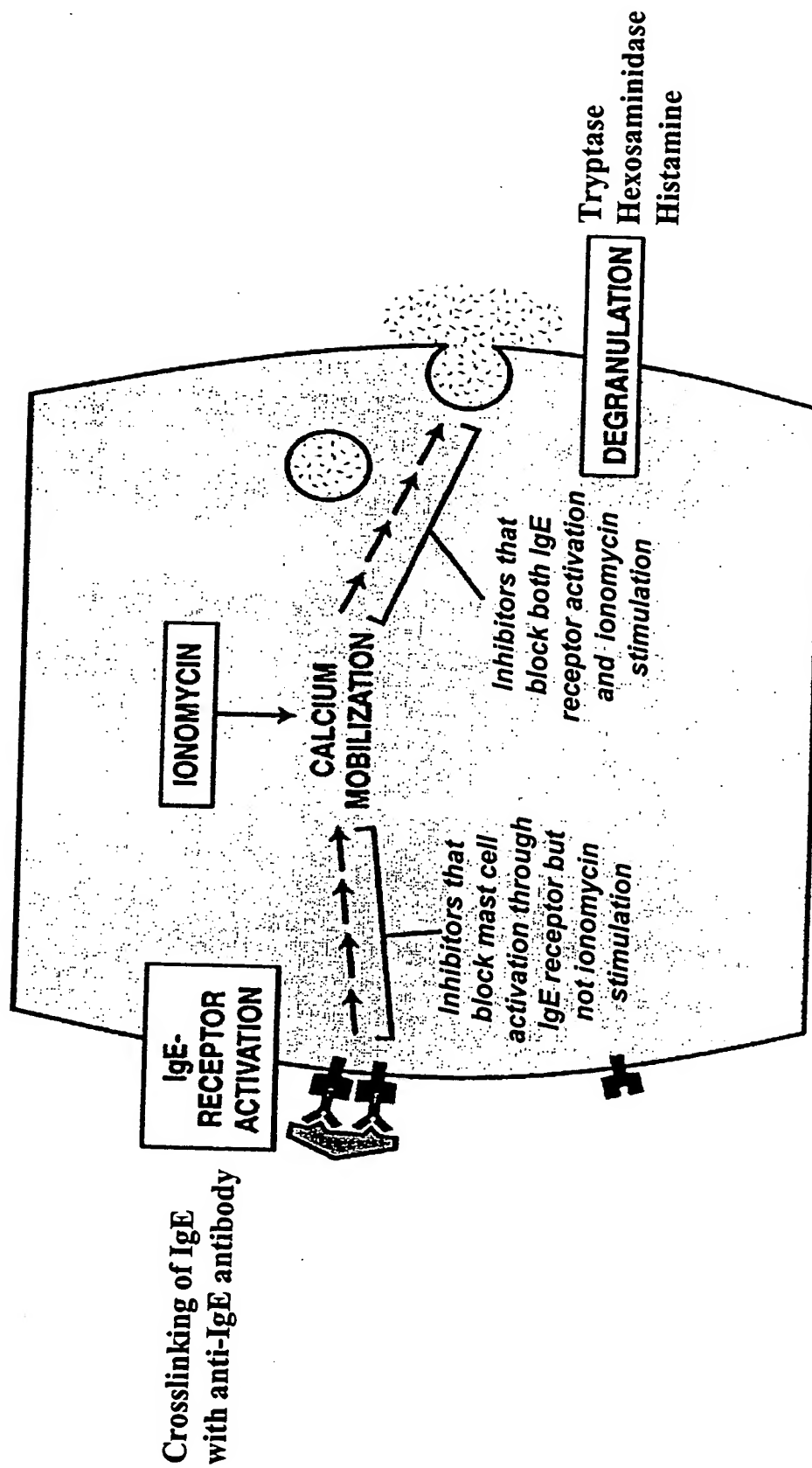


FIG. 4

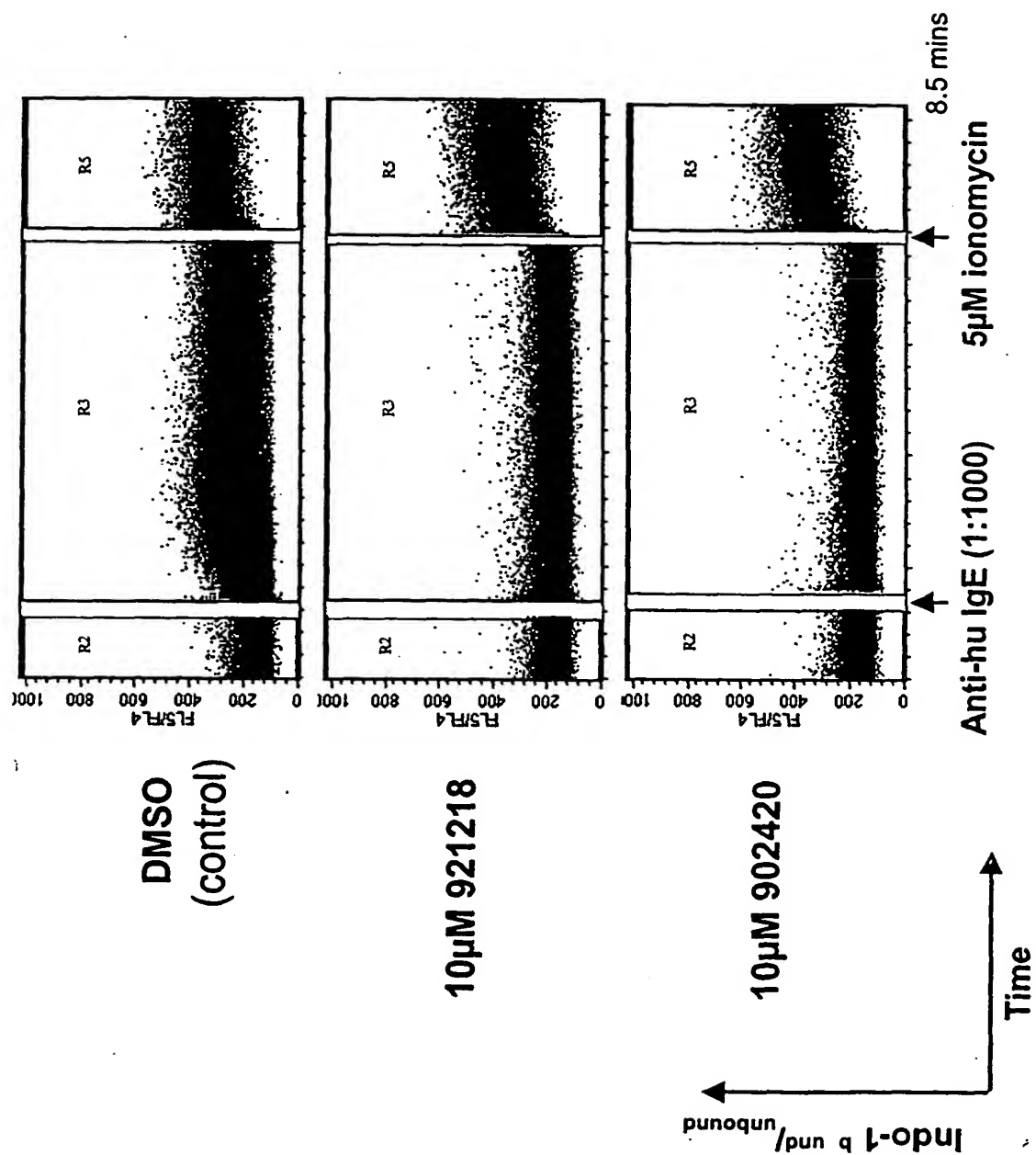


FIG. 5

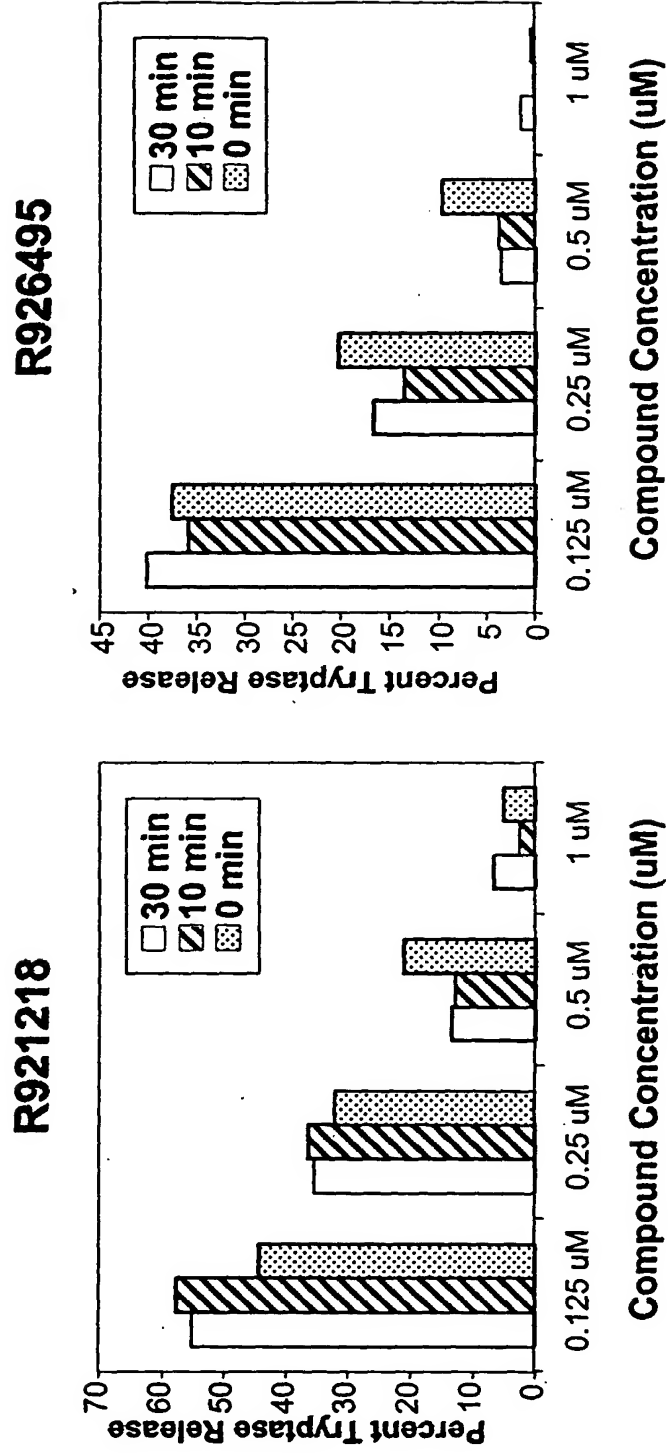


FIG. 6

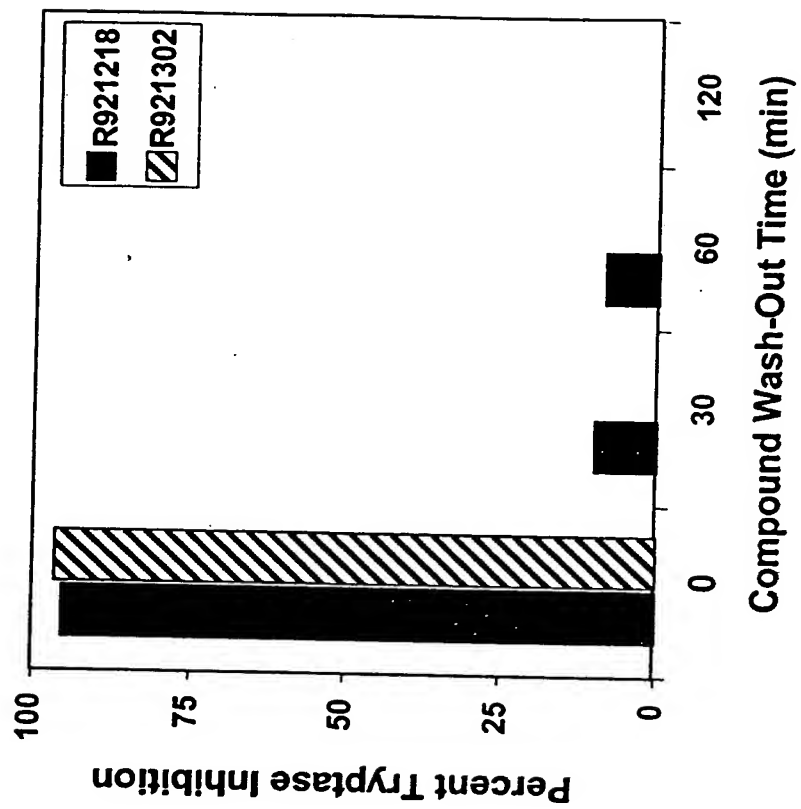


FIG. 7

Inhibition of Phosphorylation of Proteins Downstream of Syk Kinase in Fce Receptor Activated BMMC Cells

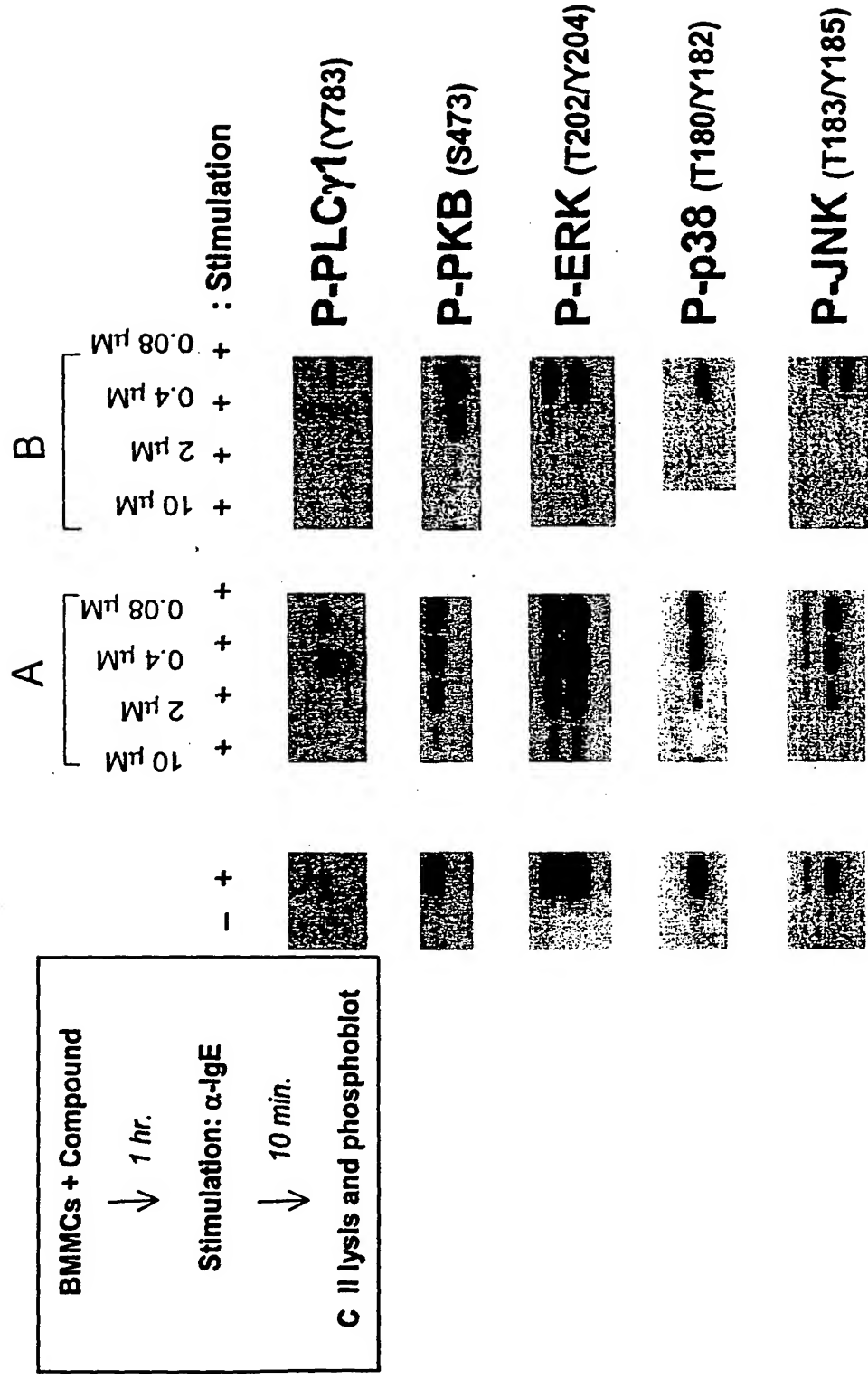
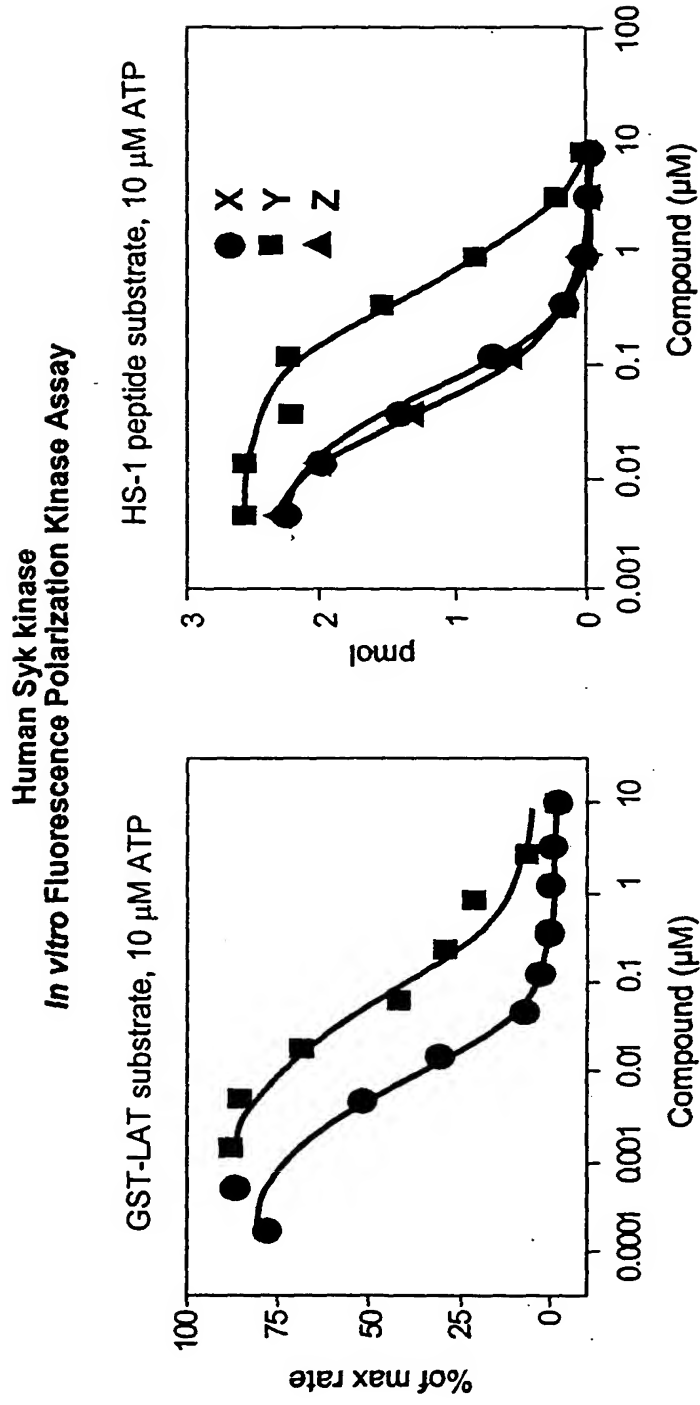


FIG. 8

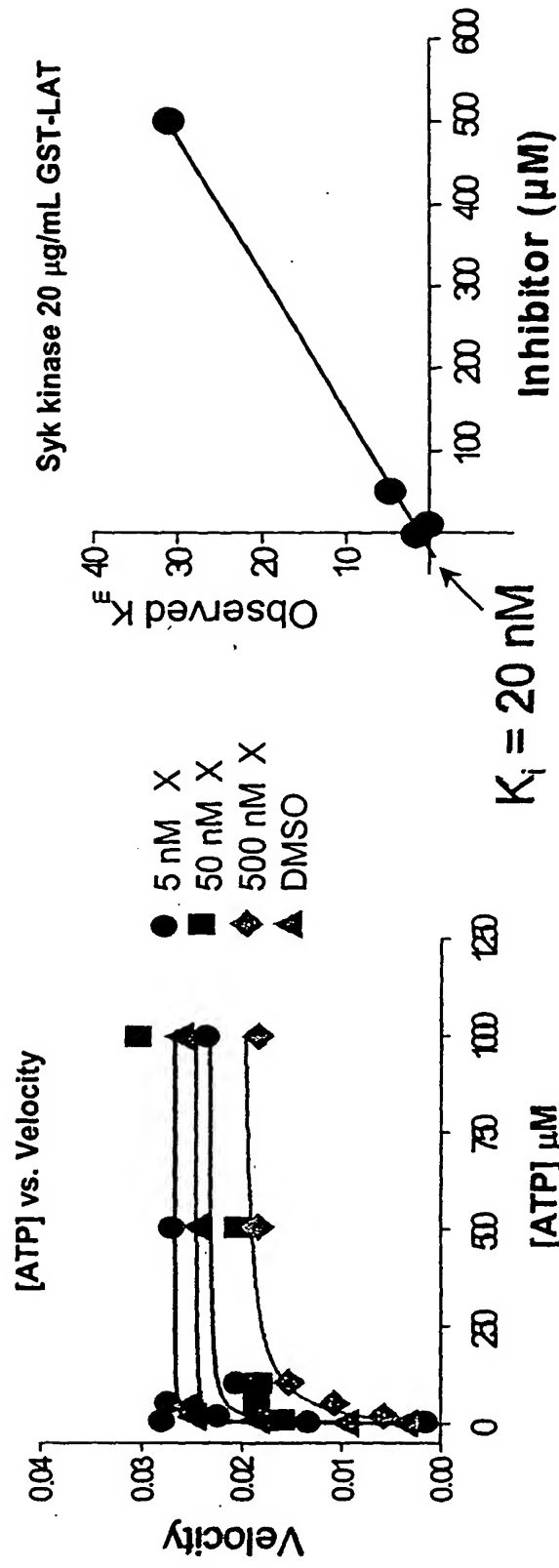
The Disclosed Compounds Potently Inhibit the Activity of Syk Kinase



IC ₅₀ (nM)	
GST-LAT	HS-1
Y	200
X	10
Z	ND
	570
	62
	43

FIG. 9

Compound Inhibition of Syk is ATP Competitive



	DMSO	5 nM X	50 nM X	500 nM X
V_{max}	.025	0.027	.023	0.020
K_m	1.54	0.79	4.5	31

CHMC: Cultured human mast cells

FIG. 10

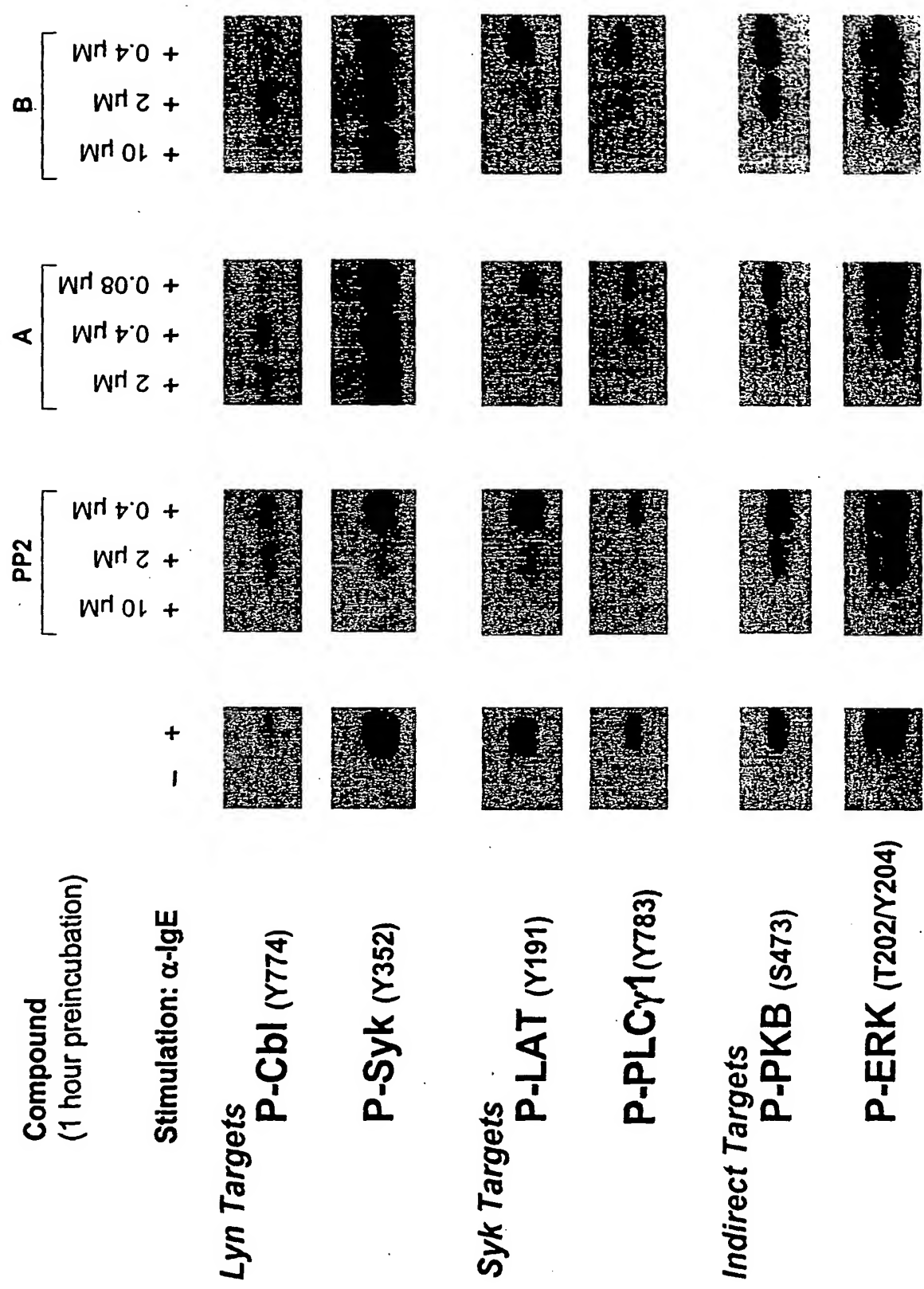


FIG. 11A

Inhibition of Phosphorylation of Proteins downstream of Syk in BMMC












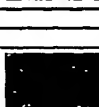





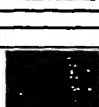


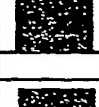
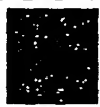





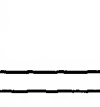

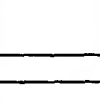


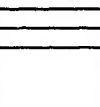





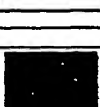
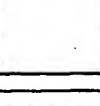

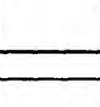


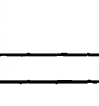


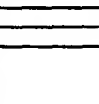












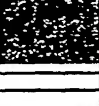
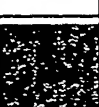







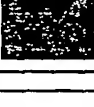

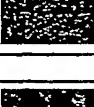


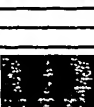
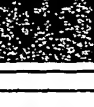

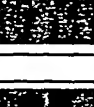
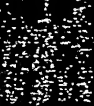





	R921219				R921304				R940323				R935138								
	+	-	+ 10 μ M	+ 2 μ M	+ 0.4 μ M	+ 0.08 μ M		+	-	+ 10 μ M	+ 2 μ M	+ 0.4 μ M	+ 0.08 μ M		+	-	+ 10 μ M	+ 2 μ M	+ 0.4 μ M	+ 0.08 μ M	
P-Syk ₃₅₂																					
P-Plc γ ₇₈₃																					
P-Lat ₁₉₁																					
P-ERK _{202/204}																					

FIG. 11B

Inhibition of Phosphorylation of Proteins downstream of Syk in BMMC

	R921303						R940347						R926891						R920410					
	-	+	10 μ M	2 μ M	0.4 μ M	+ 0.08 μ M	-	+	10 μ M	2 μ M	0.4 μ M	+ 0.08 μ M	-	+	10 μ M	2 μ M	0.4 μ M	+ 0.08 μ M	-	+	10 μ M	2 μ M	0.4 μ M	+ 0.08 μ M
P-Syk ₃₅₂																								
P-Plcγ ₇₈₃																								
P-Lat ₁₉₁																								
P-ERK _{202/204}																								

FIG. 11C

Inhibition of Phosphorylation of Proteins downstream of Syk in BMMC

	R926321				R950368				R926594				R935310			
	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+
	1	10 μ M	2 μ M	0.4 μ M	0.08 μ M	1	10 μ M	2 μ M	0.4 μ M	0.08 μ M	1	10 μ M	2 μ M	0.4 μ M	0.08 μ M	
P-Syk ₃₅₂																
P-Plc ₇₈₃																
P-Lat ₁₉₁																
P-ERK _{202/204}																

FIG. 11D

Inhibition of Phosphorylation of Proteins downstream of Syk in BMMC





























































	R935237						R926813						R926839						R908712					
	-	+	10 μ M	2 μ M	0.4 μ M	+ 0.08 μ M	-	+	10 μ M	2 μ M	0.4 μ M	+ 0.08 μ M	-	+	10 μ M	2 μ M	0.4 μ M	+ 0.08 μ M	-	+	10 μ M	2 μ M	0.4 μ M	+ 0.08 μ M
P-Syk ₃₅₂																								
P-PLC γ ₇₈₃																								
P-Lat ₁₉₁																								
P-ERK _{202/204}																								

FIG. 12

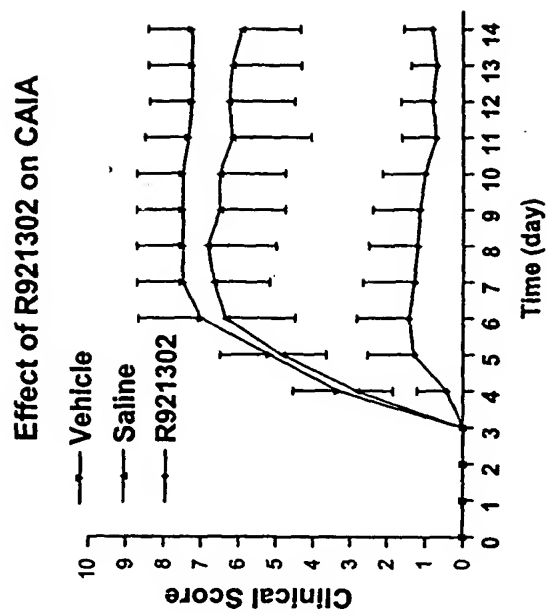


FIG. 13

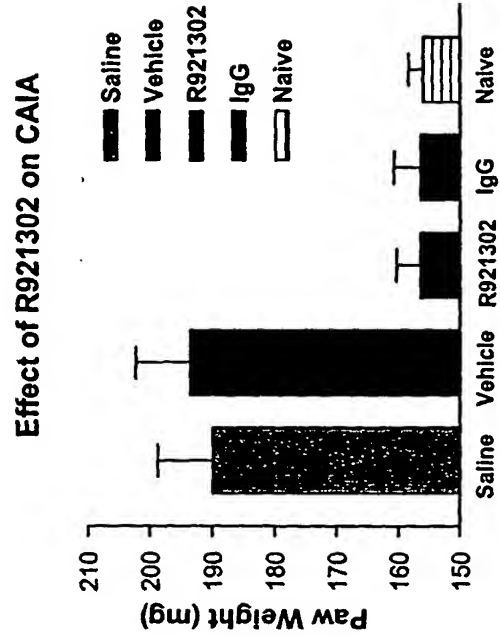


FIG. 14

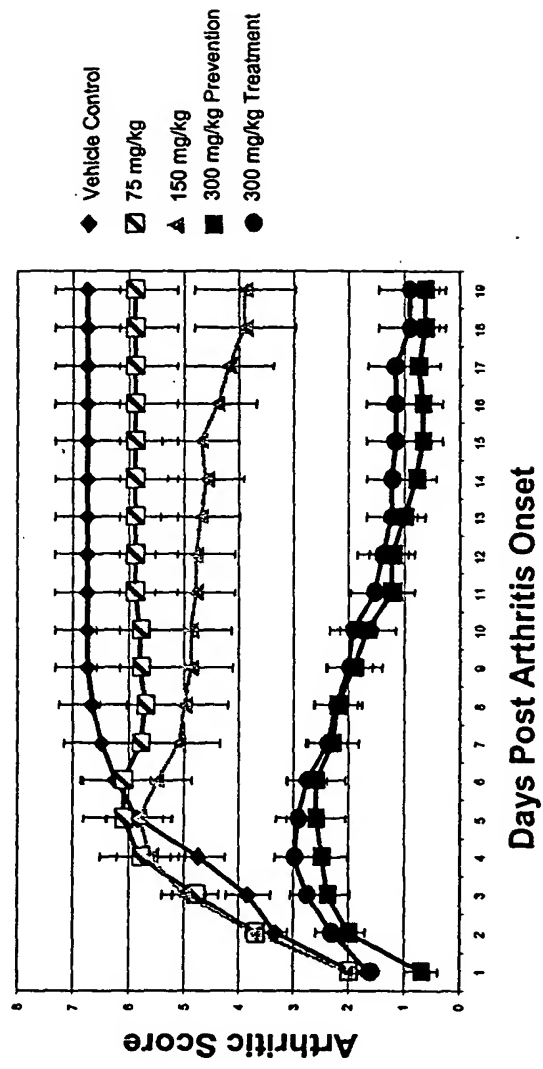


FIG. 15

Effect of R921302 on Suppression of EAE

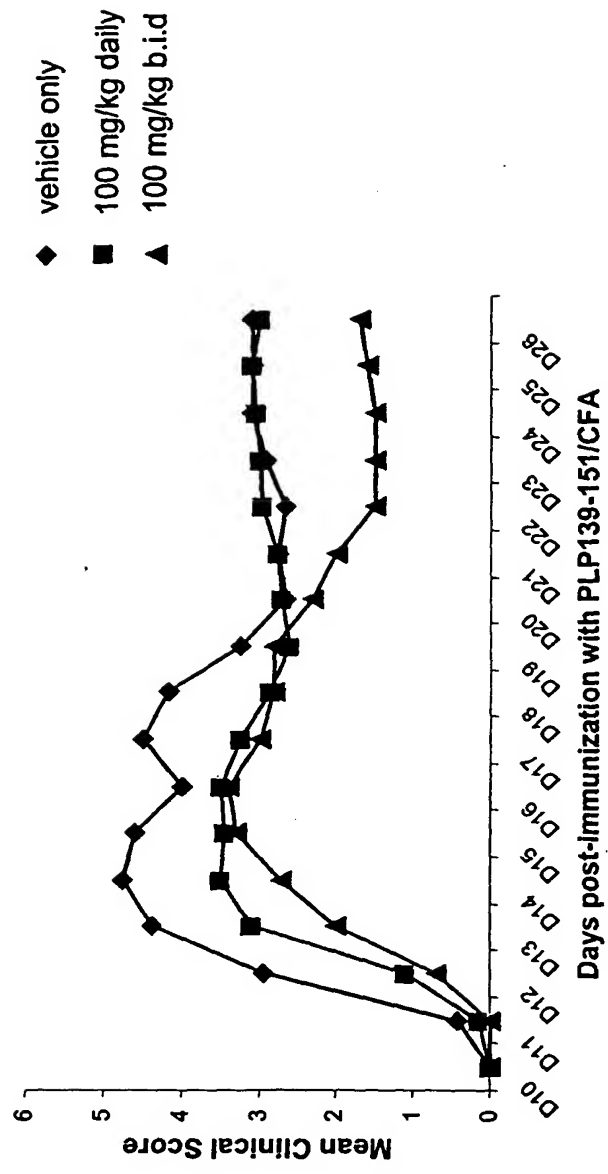


FIG. 16

